v	Mean Fluorescence Intensity	ence Intensity	% Gated	ated
mAb	L1.2-CCR5	PBMC	L1.2-CCR5	PBMC
mouse IgG1	10	4	1	-
2D7	75	29	92	36
PA8	48	6	73	3
PA9	79	\$	96	٣
PA10	08	œ	96	\$
PA11	107	∞	96	10
PA12	115	∞	96	∞
PA14	81	14	96	22

FIGURE 2

INHIBITOR	CONCENTRATION	. 65 . 77	COMBINAT	ION INDEX
COMBINATION	RATIO	ASSAY	90% Inhibition	50% Inhibition
				ļ
PA12:2D7	10:1	Entry	0.043	0.291
	2:1	Fusion	0.017	0.019
	10:1	Fusion	0.087	0.067
	50:1	Fusion	0.158	0.046
PA12:PA14	10:1	Entry	0.437	0.535
•	10:1	Fusion	0.22	0.263
		_		1.05
PA14:2D7	1:1	Entry	2.85	1.85
	1:1	Fusion	1.34	1.74
PA12:PA11	1:1	Entry	0.707	0.641
PA12:RANTES	1000:1	Fusion	0.331	0.156
PA14:RANTES	100:1	Fusion	1.6	1.37
2D7:RANTES	100:1	Fusion	0.972	0.962
PA12:CD4-IgG2	10:1	Fusion	0.3	0.31
PA14:CD4-IgG2	1:1	Fusion	0.957	0.566
2D7:CD4-IgG2	1:1	Fusion	1.127	0.302

FIGURE 3

_			IC ₅₀ valt	IC ₅₀ values (µg/ml)	
	Epitopes	cell-cell fusion inhibition	viral entry inhibition	gp120-binding inhibition	calcium flux
PA8	ž	•			
PA9	NVECL2	,		0.24	•
PA10	NvECL2			0.13	•
PA11	ž	25.5	•	0.33	•
PA12	ž	, 10.0	•	0.24	•
PA14	NVECL2	1.7	.024	1.58	6.4
2D7	ECL2	1.6	.026	1.38	45

4/20

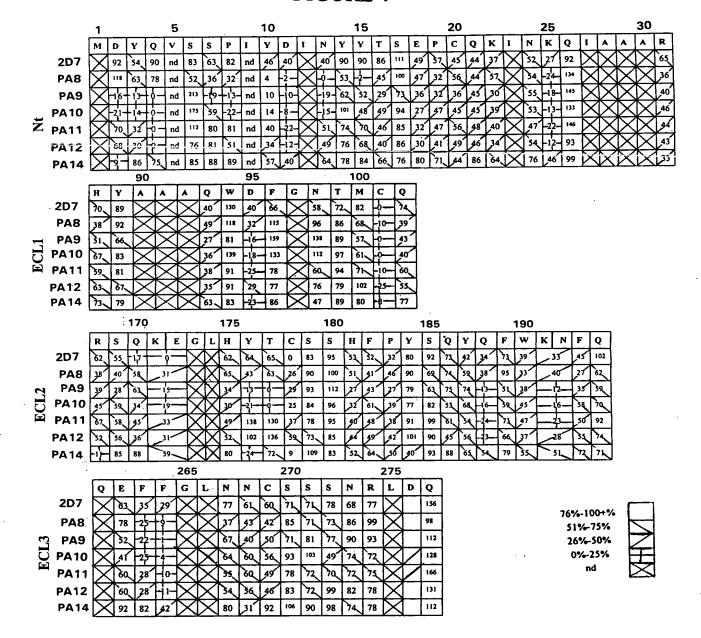


FIGURE 5A

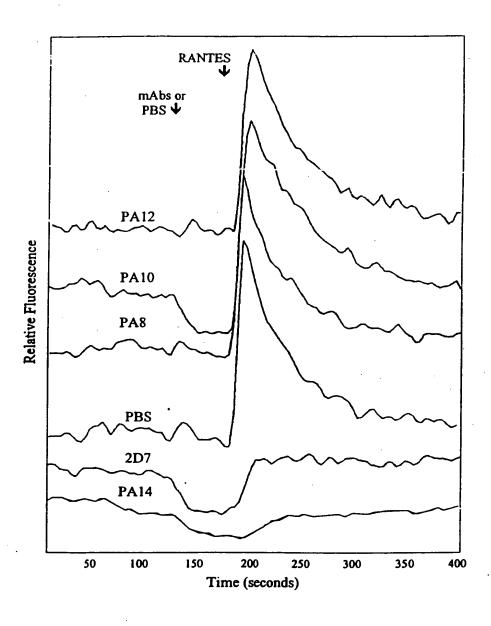


FIGURE 5B

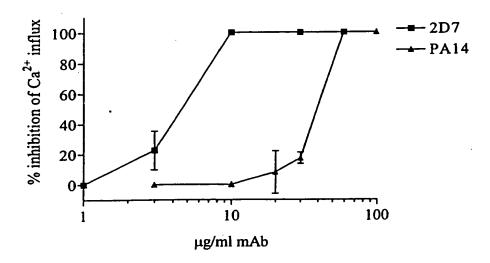


FIGURE 6A

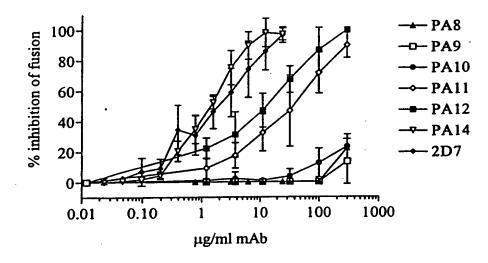


FIGURE 6B

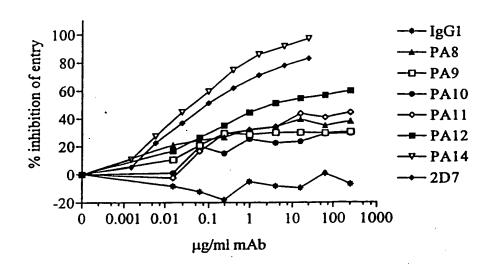


FIGURE 6C

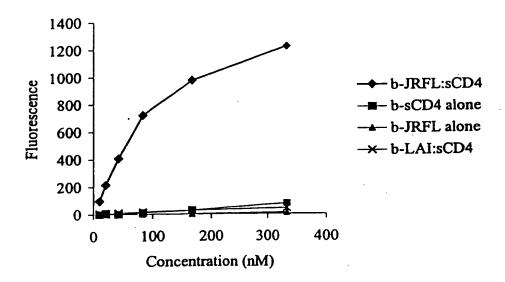
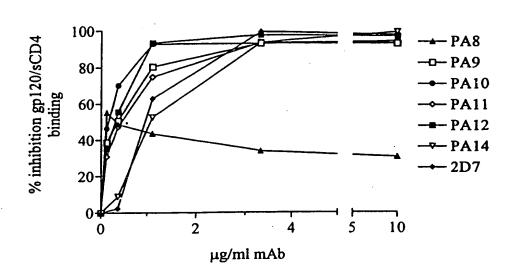
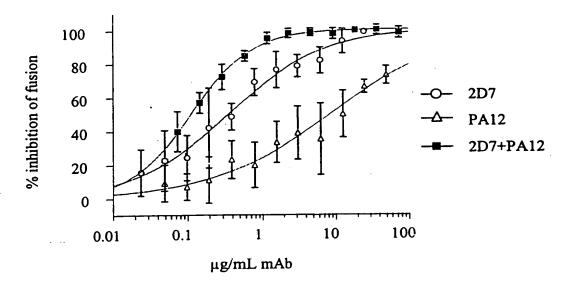
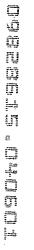
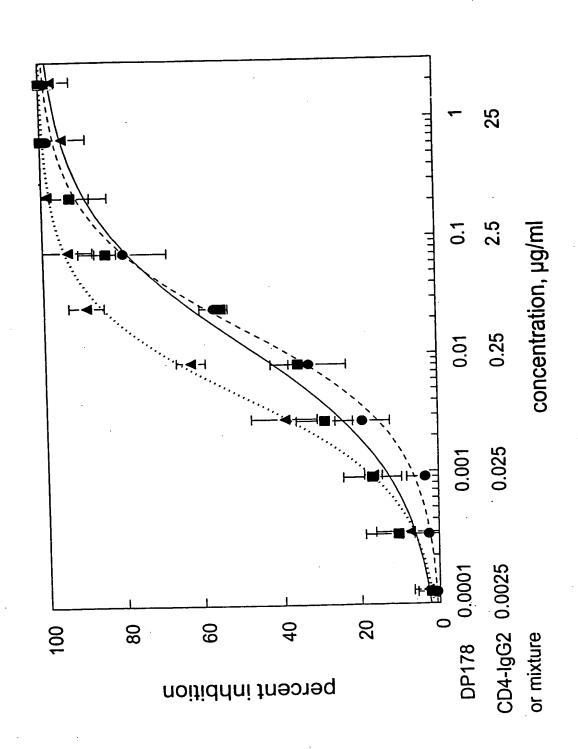


FIGURE 6D









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FIGURE 9

	С	ombinati	on Inde	X
	CD4-1	IgG2:T-2	0 Mass	Ratio
Percent Inhibition	25:1 (low)	25:1 (high)	5:1	1:1
95	0.32	0.20	0.22	0.50
90	0.38	0.25	0.27	0.55
85	0.43	0.29	0.30	0.59
80	0.47	0.33	0.34	0.62
75	0.51	0.36	0.37	0.65
70	0.54	0.39	0.40	0.67
65	0.58	0.42	0.43	0.70
60	0.61	0.45	0.45	0.73
55	0.65	0.48	0.49	0.75
50	0.69	0.51	0.52	0.78

TOSCHOT STREEDED FIGURE 10

	Dose	Reduction	29	17	13	8.4	6.3
CD4-IgG2	Concentration, µg/ml	Alone Combination	4.3	1.10	0.59	0.19	0.095
	Concent	Alone	130	19	7.8	1.6	09.0
•	Dose	Reduction	9.9	4.9	4.2	3.3	2.8
T-20	Concentration, µg/ml	Inhibition Alone Combination	0.17	0.044	0.024	0.0076	0.0039
	Concer	Alone	1.1	0.21	0.10	0.025	0.011
	Percent	Inhibition	66	95	06	70	50

FIGURE 11A

			CPS Odd	42		PA12			T-20	
	,	Concentration,	ration.		Concentration,	ration,	-	Concentration,	ration,	
4.000.00	Combination	Mu	, T	Dose	Mu	1	Dose	Mu	V	Dose
rercein Inhibition	referred Communication Index	Alone	Mix	Reduction	Alone	Mix	Reduction	Alone	Mix	Alone Mix Reduction Alone Mix Reduction Alone Mix Reduction
95	0.41	10	10 2.1	4.8	730	2.8	260	94	19	4.9
00	0.45	7.0	1.6	4.4	320	2.1	150	63	14	4.5
? ?	0.47	4.1	0.92	4.5	72	1.2	09	30	8.1	3.7
2 6	0.48	3.1	99.0	3.1 0.66 4.7	28	28 0.87	32	19	19 5.8	3.3

PRO 542, PA12 and T-20 were used in an approximate 1:1:10 molar concentration ratio.

FIGURE 11B

			PRO 542	42		PRO 140	40		T-20	
	•	Concentration.	tration.		Concentration,	tration,		Concentration,	tration,	
1	Combination	Mu	×	Dose	nM	Z	Dose	Nu I	mM	Dose
Inhibition	3	Alone	Mix	Alone Mix Reduction Alone Mix Reduction Alone Mix Reduction	Alone	Mix	Reduction	Alone	Mix	Reduction
95	0.40	8.5 1.9	1.9	4.5	19	1.0	19	140	17	8.2
90	0.39	7.1	1.5	4.7	13	0.77	17	100	13	7.7
70	0.37	5.3	0.87	6.1	7.2	0.46	16	57	7.7	7.4
20	0.35	4.6	4.6 0.63	7.3	1	4.9 0.34	14	40	5.6	7.1

PRO 542, PRO 140 and T-20 were used in an approximate 2:1:20 molar concentration ratio.

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FIGURE 11C

			PRO 542	42		PRO 140	40		T-20	0
		Concen	Concentration,		Concentration,	tration,		Concentration,	tration	
Percent	Percent Combination	[u	nM	Dose	ū	nM	Dose	Įu	nM	Dose
Inhibition	Index	Alone	Mix	Alone Mix Reduction Alone Mix	Alone	Mix	Reduction Alone Mix	Alone	Mix	Reduction
95	0.24	61	2.5	24	11.9	11.9 0.72	17	156	22	7.1
06	0.22	32	1.4	23	8.4	0.40	21	96	13	7.4
70	0.19	8.6	0.50	20	4.5	0.14	32	40	4.5	8.9
20	0.18	4.7	0.26	18	3.0	0.074	41	23	2.3	10

PRO 542, PRO 140 and T-20 were used in an approximate 4:1:30 molar concentration ratio.

int Co	,						
	in officer	Concentration,	tration,		Concentration,	tration,	
,		Mu	7	Dose	nM	¥	D)ose
ļ	Index	Alone Mix	1 1	Reduction Alone Mix	Alone		Reduction
 	0.56	12	1.8	6.7	156	55	2.8
06	0.55	8.4	1.1	7.4	96	35	2.7
70 07	0.55	4.5	0.51	8.8	40	16	2.5
. 0 0.	0.56	3.0	3.0 0.31	6.6	23	10	2.4

PRO 140 and T-20 were used in an approximate 1:30 molar concentration ratio.

TOYONE 12 TENEND

Triple Combination Synergistically Blocks HIV-1 Entry (I)

